PTO/SB/21 (08-00) Cacy e

(L)			Application Numb r	ed for use through 10/31/2002. OMB 0651 rk Office: U.S. DEPARTMENT OF COMMI ormation unless it displays a valid OMB control no		
A IRANSWIII IAL			Filing Date	May 12, 1998		
FOI	RM		First Named Inventor	David J. Ecker		
(to be used for all correspo	ondence after i	nitial filing)	Group Art Unit			
			Examiner Name			
Total Number of Pages in T	his Submission		Attorney Docket Number	IBIS0040-100/IBIS-0007		
		ENCL	OSURES (check all that apply)			
Fee Transmittal Form			ment Papers Application)	After Allowance Communication Group		
Fee Attached		☐ Drawin	g(s)	Appeal Communication to Board Appeals and Interferences		
Amendment / Respons	е	Licensi	ing-related Papers	Appeal Communication to Group (Appeal Notice, Brief, Reply Brief)		
After Final		Petition	n	Proprietary Information		
Affidavits/declaration	on(s)		n to Convert to a ional Application	Status Letter		
Extension of Time Req	uest		of Attorney with Revocation nange of Correspondence ss	Other Enclosure(s) (please identify below):		
Express Abandonment	Request	1 =	al Disclaimer st for Refund	Table A Statement Under 37 CFR 3.7 with executed Assignment		
☐ Information Disclosure Statement			umber of CD(s)	with excedice Assignment		
Certified Copy of Priori	ty	Rema				
Response to Missing P						
Response to Missir Parts under 37 CF 1.52 or 1.53						
	SIGNA	TURE OF	APPLICANT, ATTORNEY,	OR AGENT		
Firm or Paul K. Legaard, Registration No. 38,534 Individual name						
Signature Pellfy Date 23 DECEMBER 2003						
Date 23	DECEM	BER 2	000			
			RTIFICATE OF MAILING			
			sited with the United States Posashington, D.C. 20231 on this d	tal Service as first class mail in an enve ate: [2] DEUMBER 2007]		
	, , , , , , , , , , , , ,			- UUGITIOU C MO J		
Typed or printed name	Paul K. Lega	ard, Registra	tion No. 38,534			

Burden Hour Statement: This form is estimated to take 0.2 hours to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be send to the Chief Information Officer, U.S. Patent and Trademark Office, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Assistant Commissioner for Patents, Washington, DC 20231.



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Assistant Commissioner for Patents Washington, DC 20231

Sir:

POWER OF ATTORNEY WITH REVOCATION AND CHANGE OF CORRESPONDENCE ADDRESS

I hereby revoke all previous powers of attorney or authorizations of agent given in the patents/applications identified in Table A attached hereto, and appoint

patents/applications identified in Table A attached hereto, and appoint
Practitioners at Customer Number: 34138 as my/our attorney(s) or agent(s) to prosecute the patents/applications identified in Table A, and to transact all business in the United States Patent and Trademark Office connected therewith.
Please change the Correspondence Address for the patent applications identified in
Table A to:
Practitioners at Customer Number: 34138
I am the: Applicant/Inventor. Assignee of record of the entire interest. (A statement under 37 CFR 3.73(b) is enclosed.)
Respectfully submitted,
ISIS PHARMACEUTICALS, INC.
Date: 16-63 By: Aname: B. Lynne Parshall Title: Executive Vice President



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Assistant Commissioner for Patents Washington, DC 20231

Sir:

STATEMENT UNDER 37 CFR 3.73(b)

Applicant, ISIS Pharmaceuticals, Inc., a corporation, states that it is:

the assignee of the entire right, title, and interest of the patent applications identified in Table A attached hereto.

The extent (by, percentage) of its ownership interest is 100% in the patent applications identified in Table A.

a copy of the pertinent assignment from the inventor(s) of the patent applications identified in Table A is enclosed herewith.

The undersigned (whose title is supplied below) is authorized to act on behalf of the assignee.

Respectfully submitted,

ISIS PHARMACEUTICALS, INC.

Date: 11-6-03

By:

Name: B. Lynne Parshall

Title: Executive Vice President

18150040-100

PATENT Joint Inventors

(Pending Application; Serial No. 09/076,404)

DOCKET NO.: IBIS-0007

ASSIGNMENT

WHEREAS, we David J. Ecker, Richard Griffey, Stanley T. Crooke, Ranga Sampath, Eric Swayze, Venkatraman Mohan and Stev Hofstadler hereinafter referred to as the assignors, residing respectively at 1041 Saxony Road, Encinitas, CA 92024, 360 Barsby Street, Vista, CA 92084, 3211 Piragua Street, Carlsbad, CA 92009, 12223 Mannix Road, San Diego, CA 92129, 7789 Palenque Street Carlsbad, CA 92009, 7042 Via Cabana, Carlsbad, CA 92009 and 5014 Viewridge Way, Oceanside, CA 92056 are the joint inventors of certain inventions or improvements for which we have made application for Letters Patent to the United States, identified as Case No. IBIS-0007, entitled MODULATION OF MOLECULAR INTERACTION SITES ON RNA AND OTHER BIOMOLECULES and

WHEREAS, ISIS Pharmaceuticals, Inc. hereinafter referred to as the assignee, of 2292 Faraday Avenue, Carlsbad CA 92008, a corporation of Delaware, is desirous of acquiring the entire right, title and interest in and to the said inventions or improvements and in and to the said application, and in, to and under any and all Letters Patent which may be granted on or as a result thereof in any and all countries:

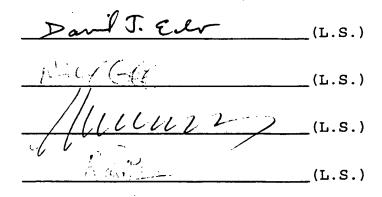
NOW, THEREFORE, for and in consideration of the sum of One Dollar (\$1.00) to each of us in hand paid by said assignee, and other good and valuable consideration, the receipt of which is hereby acknowledged, we, the said assignors, have sold, assigned, transferred and set over, and by these presents do hereby sell, assign, transfer and set over to said assignee, the entire right, title and interest in and to said inventions or improvements and said application and any and all continuations, divisions and renewals of and substitutes for said application, and in, to and under any and all Letters Patent which may be granted on or as a result thereof in the United States and any and all other countries, and any reissue or reissues or extension or extensions of said Letters Patent, and assign to and authorize said assignee, to file in our names applications for Letters Patent in all countries, the same to be held and enjoyed by said assignee, its successors, assigns, nominees or legal representatives, to the full end of the term or terms for which said Letters Patent respectively may be granted, reissued or extended, as fully and entirely as the same would have been held and enjoyed by us had this assignment, sale and transfer not been made.

AND we hereby covenant that we have full right to convey the entire interest herein assigned, and that we have not executed and will not execute any agreement in conflict herewith, and we further covenant and agree that we will each time request is made and without undue delay, execute and deliver all such papers as may be necessary or desirable to perfect the title to said inventions or improvements, said application and said Letters Patent to said assignee, its successors, assigns, nominees, or legal representatives, and each of us agrees to communicate to said assignee or to its nominee all known facts respecting said inventions or improvements, said application and said Letters Patent, to testify in any legal proceedings, to sign all lawful papers, to execute all disclaimers and divisional, continuing, reissue and foreign applications, to make all rightful oaths, and generally to do everything possible to aid said assignee, its successors, assigns, nominees and legal representatives to obtain and enforce for its or their own benefit proper patent protection for said inventions or improvements in any and all countries.

AND we hereby authorize and request the Commissioner of Patents and Trademarks of the United States and any official of any country or countries foreign to the United States whose duty it is to issue patents on applications as aforesaid, to issue to said assignee, as assignee of the entire right, title and interest, any and all Letters Patent for said inventions or improvements, including any and all Letters Patent of the United States which may be issued and granted on or as a result of the application aforesaid, in accordance with the terms of this assignment.

We further authorize and direct our attorneys to insert below* the serial number and filing date of said application now identified as Case No. IBIS-0007 as soon as the same shall have been made known to them by the United States Patent Office.

IN WITNESS WHEREOF, we have hereunto set our hands and seals.



PATENT Joint Inventors

Elin Eswyt	(L.s.)
Undan	(L.S.)
Steve A. Agtable	(L.S.
	(T. S.

[WHERE ALL INVENTORS SIGN BEFORE THE SAME NOTARY:]
CALIFCENIA:
CALIFCRNIA: SS COUNTY OF SANDIEGE:
On this 26th day of MAY, 1998, before me
STANLEY T. CROOKE, RANGA SAMPATH ERIC JWAYZE, VENKATRAMAN MOHAN, STEVE HOFSTADLER to me personally known and known to me to be the same individuals who executed the foregoing assignment, and who acknowledged to me that they executed the same of their own free will for the use and purposes therein set forth.
Notary Public
Notary Public
[WHERE INVENTORS SIGN BEFORE SEPARATE NOTARIES:]::::
COUNTY OF:
On this day of, 19, before me personally came the above named to me personally known and known to me to be the same individual who executed the foregoing assignment, and who acknowledged to me that execution of the same was of that person's own free will for the use and purposes therein set forth.
Notary Public

Table A

١٩٠١		<u> </u>			I	<u> </u>		т
Title Title	Identification of Disease Predictive Nucleic Acids	IN SITU Binary Synthesis of Biologic00ally Effective Molecules	IN SITU Binary Synthesis of Biologically Effective Molecules	Identification of Molecular Interaction Sites in RNA for Novel Drug Discovery	Identification of Molecular Interaction Sites in RNA for Novel Drug Discovery	Generation of Virtual Combinatorial Libraries of Compounds	Generation of Virtual Combinatorial Libraries of Compounds	Characterization of Interactions Between Molecular Interaction Sites of RNA and Ligands Therefor
Art Unit	1655	1634	1634	1656	1634	2123	1631	1655
Examiner	J. Fredman	J. Fredman	To Be Assigned	E. Campbell	F. Lu	S. Broda	M. Borin	J. Fredman
Inventor(s)	David J. Ecker	David J. Ecker	David J. Ecker	David J. Ecker et al.	David J. Ecker et al.	Richard Griffey et al.	Richard Griffey et al.	Richard Griffey et al.
Filing Date Issue Date	November 25, 1998 September 17, 2002	November 25, 1998 December 10, 1998	August 1, 2003	May 12, 1998 April 24, 2001	May 12, 1999	May 12, 1998	January 3, 2001	May 12, 1998
Serial No. Patent No. Docket No.	09/200,355 6,451,524 IBIS0010-100	09/200,107 6,492,111 IBIS0034-100	10/209,761 IBIS0034-101	09/076,440 6,221,587 IBIS0035-100	09/310,667 IBIS0035-101	09/076,405 IBIS0036-100	09/753,869 IBIS0036-101	09/076,447 IBIS0037-100

OTP &	VC/124									
NEC 2 9 2003	Characterization of Interactions Between Molecular Interaction Sites of RNA and Ligands Therefor	Mass Spectrometric Methods for Biomolecular Screening	Mass Spectrometric Methods for Biomolecular Screening	Mass Spectrometric Methods for Biomolecular Screening	Mass Spectrometric Methods for Biomolecular Screening	Generation of Combinatorial Libraries of Compounds Corresponding to Virtual Libraries of Compounds	Generation of Combinatorial Libraries of Compounds Corresponding to Virtual Libraries of Compounds	Small Molecule Targeting Of RNA Structures	Identification of Molecular Interaction Sites in RNA for Novel Drug Discovery	Modulation of Molecular Interaction Sites on RNA and Other Biomolecules
	1637	1634	1634	1634		1639	1639	Not applicable	1631	1631
	J. Fredman	S. Zitomer	S. Zitomer	A. Chakrabarti	To Be Determined	T. Friend	M. Tran	Not applicable	A. Marschel	A. Marschel
•	Richard Griffey et al.	Stanley T. Crooke et al.	Stanley T. Crooke	Stanley T. Crooke	Stanley T. Crooke	Richard H. Griffey et al.	Richard H. Griffey et al.	Stanley T. Crooke	David J. Ecker et al.	David J. Ecker et al.
	May 12, 1999	May 12, 1998 August 6, 2001	March 2, 1999 December 11, 2001	June 19, 2001	June 27, 2003	May 12, 1998	September 21, 1999	May 12, 1998	May 12, 1998	May 12, 1999
	09/310,653 IBIS0037-101	09/076,206 6,428,956 IBIS0038-100	09/260,310 6,329,146 IBIS0038-101	09/884,317 IBIS0038-102	09/884,317 IBIS0038-103	09/076,214 IBIS0039-100	09/400,150 IBIS0039-101	60/085,092 IBIS0040-001	09/076,404 IBIS0040-100	09/310,735 IBIS0040-101

O P & Walter									
LIEP F	2003								
SPANT & 12	Modulation of Molecular Interaction Sites on RNA and Grace Other Biomolecules	Molecular Interaction Sites of Vimentin RNA and Methods of Modulating the Same	Molecular Interaction Sites of Vimentin RNA and Methods of Modulating the Same	Molecular Interaction Sites of Ornithine Decarboxylase RNA and Methods of Modulating the Same	Molecular Interaction Sites of Hepatitis C Virus RNA and Methods of Modulating the Same	Methods for Identifying Ligand Binding Sites in a Biomolecule	Molecular Interaction Sites of 23S Ribosomal RNA and Methods of Modulating the Same	Molecular Interaction Sites in 23S Ribosomal RNA and Methods of Modulating the Same	Molecular Interaction Sites of RNAse P RNA and Methods of Modulating the Same
	1631	1631	1631	1635	Not Applicable	1645	Not Applicable	1645	Not Applicable
	A. Marschel	A. Marschel	A. Marschel	M. Schmidt	Not Applicable	To Be Assigned	Not Applicable	To Be Assigned	Not Applicable
•	David J. Ecker et al.	David J. Ecker et al.	David J. Ecker et al.	David J. Ecker et al.	David J. Ecker	David J. Ecker	David J. Ecker	David J. Ecker	David J. Ecker
	March 22, 2002	May 12, 1999	April 24, 2002	May 12, 1999	August 22, 2001	August 19, 2002	August 22, 2001	August 20, 2002	August 21, 2001
	10/104,949 IBIS0040-102	09/310,907 IBIS0041-100	10/135,017 IBIS0041-101	09/310,761 IBIS0042-100	60/314,236 IBIS0043-001	10/225,501 IBIS0043-100	60/314,251 IBIS0044-001	10/224,836 IBIS0044-100	60/314,020 IBIS0045-001

292 3 1920		Molecular Interaction Sites in 16S Ribosomal RNA and Methods of Modulating the Same	Molecular Interaction Sites in 16S Ribosomal RNA and Methods of Modulating the Same	Methods for Identifying Ligand Binding Sites in a Biomolecule	Methods for Identifying Ligand Binding Sites in a Biomolecule	Optimization Of Ligand Affinity For RNA Targets Using Mass Spectrometry	Optimization of Ligand Affinity For RNA Targets Using Mass Spectrometry	l Biaryl	ounds and al Libraries Same	Multi-Paradigm Knowledge- Bases
	Molecular Interaction RNAse P RNA and M Modulating the Same	Molecular In 16S Ribosor Methods of N Same	Molecular In 16S Ribosor Methods of In Same	Methods for Binding Sites	Methods for Binding Site:	Optimization (For RNA Targ Spectrometry	Optimization (For RNA Targ	Antimicrobial Biaryl Compounds	Amine Compounds and Combinatorial Libraries Comprising Same	Multi-Paradiç Bases
	1645	Not Applicable	1645	1637	1645	To Be Assigned	To Be Assigned	1624	1627	Not Applicable
	To Be Assigned	Not Applicable	To Be Assigned	J. Riley	To Be Assigned	To Be Assigned	To Be Assigned	B. Coleman	T. Prosthofer	Not Applicable
	David J. Ecker	David J. Ecker	David J. Ecker	Richard H. Griffey et al.	Richard H. Griffey et al.	Richard H. Griffey et.al.	Richard H. Griffey et al.	Elizabeth Jefferson	Eric Edward Swayze et al.	John McNeil
	August 16, 2002	August 21, 2001	August 16, 2002	February 22, 2001	July 31, 2002	February 8, 2000	May 16, 2000	August 1, 2000	September 5, 1997	May 16, 2001
	10/223,156 IBIS0045-100	60/313,890 IBIS0046-001	10/223,126 IBIS0046-100	09/791,147 IBIS0047-100	10/209,692 IBIS0047-101	09/499,875 IBIS0048-100	09/573,479 IBIS0048-101	09/630,122 IBIS0050-100	08/924,851 IBIS0051-100	60/291,459 IBIS0052-001

• -	7003		7				ı 	1		
\$15M7 & 150	Multi-Paradigm Knowledge	System and Method for Sample Positioning in a Robotic System	System and Method for Sample Positioning in a Robotic System	Novel Pyranosyl Cytosines: Pharmaceutical Formulation and Methods	Novel Benzimidazole Compounds	Self-Dispensing Storage Device	Molecular Interaction Sites of Interleukin-2 RNA and Methods of Modulating the Same	Optimization of Ligand Affinity for RNA Targets Using Mass Spectrometry	Optimization of Ligand Affinity for RNA Targets Using Mass Spectrometry	Antisense Modulation of Human MDM2 Expression
	2176	1743	1743	1624	1624	1743	1635	1639	1627	1635
	To Be Assigned	Patricia A. Bex	To Be Assigned	T. McKenzie	K. Habte	E. Quan	M. Schmidt	M. Tran	T. Prosthofer	T. Gibbs
	John McNeil et al.	John McNeil	John McNeil	Michael L. Migawa	Eric E. Swayze et al.	John McNeil	David J. Ecker et al.	Richard Griffey et al.	Richard Griffey et al.	Loren J. Miraglia et al.
	May 16, 2002	October 1, 1999 August 6, 2002	May 22, 2002	February 24, 2002	February 6, 2002	October 2, 2000	May 12, 1999	February 8, 2000	May 16, 2000	December 4, 2001
	10/150,668 IBIS0052-100	09/411,748 6,429,016 IBIS0053-100	10/153,327 IBIS0053-101	10/067,017 IBIS0054-100	10/071,978 IBIS0055-100	09/678,434 IBIS0056-100	09/310,844 IBIS0058-100	09/499,875 IBIS0059-100	09/573,479 IBIS0059-101	10/005,344 ISIS0052-100

3 5003 SE	Γ
Oligoribonucleotides and Ribonucleases for Cleaving	Antisense Modulation of SMAD6 Expression
1635	1635
To Be Assigned	To Be Assigned
Stanley T. Crooke	Brett P. Monia et al.
October 25, 2002	December 20, 2002
10/281,349 ISIS0002-105	10/327,805 ISIS0060-101